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&lt;110&gt; Zimmerman, et al.

&lt;120&gt; M-CSF MUTEINS AND USES THEREOF

&lt;130&gt; PP021603.0002

&lt;140&gt; To be assigned

&lt;141&gt; 2005-01-21

&lt;150&gt; US 60/537,985

&lt;151&gt; 2004-01-21

&lt;160&gt; 8

&lt;170&gt; PatentIn version 3.2

&lt;210&gt; 1

&lt;211&gt; 1855

&lt;212&gt; DNA

&lt;213&gt; Homo sapiens

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&lt;222&gt; (286)..(957)

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Tyr Leu Lys Lys Ala Phe Leu Leu Val Gln Asp Ile Met Glu Asp Thr			
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165

170

175

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&lt;213&gt; Homo sapiens

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Tyr Leu Lys Lys Ala Phe Leu Leu Val Gln Asp Ile Met Glu Asp Thr			
85	90	95	

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Gln Pro Leu Ala Pro Ser Met Ala Pro Val Ala Gly Leu Thr Trp Glu  
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Pro Glu Glu Ala Ser Gly Glu Ala Ser Glu Ile Pro Val Pro Gln Gly  
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Thr Glu Leu Ser Pro Ser Arg Pro Gly Gly Ser Met Gln Thr Glu  
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Ser Val Leu Pro Leu Gly Glu Leu Glu Gly Arg Arg Ser Thr Arg Asp  
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Ala Arg Pro Leu Pro Arg Phe Asn Ser Val Pro Leu Thr Asp Thr Gly  
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His Glu Arg Gln Ser Glu Gly Ser Ser Ser Pro Gln Leu Gln Glu Ser  
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Val Phe His Leu Leu Val Pro Ser Val Ile Leu Val Leu Leu Ala Val  
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Gly Gly Leu Leu Phe Tyr Arg Trp Arg Arg Arg Ser His Gln Glu Pro  
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Cys Gln Ile Thr Phe Glu Phe Val Asp Gln Glu Gln Lys Asp Pro				
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ctc ttg ttc tac agg tgg agg ccg ccg agc cat caa gag cct cag aga Leu Leu Phe Tyr Arg Trp Arg Arg Arg Ser His Gln Glu Pro Gln Arg 400 405 410			1431
gcg gat tct ccc ttg gag caa cca gag ggc agc ccc ctg act cag gat Ala Asp Ser Pro Leu Glu Gln Pro Glu Gly Ser Pro Leu Thr Gln Asp 415 420 425 430			1479
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Gln Ser Leu Gln Arg Leu Ile Asp Ser Gln Met Glu Thr Ser Cys Gln  
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Ile Thr Phe Glu Phe Val Asp Gln Glu Leu Lys Asp Pro Val Cys  
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Tyr Leu Lys Lys Ala Phe Leu Leu Val Gln Asp Ile Met Glu Asp Thr  
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Met Arg Phe Arg Asp Asn Thr Pro Asn Ala Ile Ala Ile Val Gln Leu  
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Gln Glu Leu Ser Leu Arg Leu Lys Ser Cys Phe Thr Lys Asp Tyr Glu  
115 120 125

Glu His Asp Lys Ala Cys Val Arg Thr Phe Tyr Glu Thr Pro Leu Gln  
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Leu Leu Glu Lys Val Lys Asn Val Phe Asn Glu Thr Lys Asn Leu Leu  
145 150 155 160

Asp Lys Asp Trp Asn Ile Phe Ser Lys Asn Cys Asn Asn Ser Phe Ala  
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Glu Cys Ser Ser Gln Asp Val Val Thr Lys Pro Asp Cys Asn Cys Leu  
180 185 190

Tyr Pro Lys Ala Ile Pro Ser Ser Asp Pro Ala Ser Val Ser Pro His  
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Gln Pro Leu Ala Pro Ser Met Ala Pro Val Ala Gly Leu Thr Trp Glu  
210 215 220

Asp Ser Glu Gly Thr Glu Gly Ser Ser Leu Leu Pro Gly Glu Gln Pro  
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Leu His Thr Val Asp Pro Gly Ser Ala Lys Gln Arg Pro Pro Arg Ser

245

250

255

Thr Cys Gln Ser Phe Glu Pro Pro Glu Thr Pro Val Val Lys Asp Ser  
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Thr Ile Gly Gly Ser Pro Gln Pro Arg Pro Ser Val Gly Ala Phe Asn  
 275 280 285

Pro Gly Met Glu Asp Ile Leu Asp Ser Ala Met Gly Thr Asn Trp Val  
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Pro Glu Glu Ala Ser Gly Glu Ala Ser Glu Ile Pro Val Pro Gln Gly  
 305 310 315 320

Thr Glu Leu Ser Pro Ser Arg Pro Gly Gly Ser Met Gln Thr Glu  
 325 330 335

Pro Ala Arg Pro Ser Asn Phe Leu Ser Ala Ser Ser Pro Leu Pro Ala  
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Ser Ala Lys Gly Gln Gln Pro Ala Asp Val Thr Gly His Glu Arg Gln  
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Ser Glu Gly Ser Ser Ser Pro Gln Leu Gln Glu Ser Val Phe His Leu  
 370 375 380

Leu Val Pro Ser Val Ile Leu Val Leu Leu Ala Val Gly Gly Leu Leu  
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Phe Tyr Arg Trp Arg Arg Arg Ser His Gln Glu Pro Gln Arg Ala Asp  
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gccctggca ctttcagcc ctgcacactgc ctgccacttc cccaccgagg cc atg ggc      298
                                         Met Gly
                                         1

cca gga gtt ctg ctg ctc ctg ctg gtc aca gct tgg cat ggt cag      346
Pro Gly Val Leu Leu Leu Leu Leu Val Ala Thr Ala Trp His Gly Gln
      5           10          15

gga atc cca gtg ata gag ccc agt gtc cct gag ctg gtc gtg aag cca      394
Gly Ile Pro Val Ile Glu Pro Ser Val Pro Glu Leu Val Val Lys Pro
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Gly Ala Thr Val Thr Leu Arg Cys Val Gly Asn Gly Ser Val Glu Trp
      35          40          45          50

gat ggc ccc cca tca cct cac tgg acc ctg tac tct gat ggc tcc agc      490
Asp Gly Pro Pro Ser Pro His Trp Thr Leu Tyr Ser Asp Gly Ser Ser
      55          60          65

agc atc ctc agc acc aac aac gct acc ttc caa aac acg ggg acc tat      538
Ser Ile Leu Ser Thr Asn Asn Ala Thr Phe Gln Asn Thr Gly Thr Tyr
      70          75          80

cgc tgc act gag cct gga gac ccc ctg gga ggc agc gcc gcc atc cac      586
Arg Cys Thr Glu Pro Gly Asp Pro Leu Gly Gly Ser Ala Ala Ile His
      85          90          95

ctc tat gtc aaa gac cct gcc cgg ccc tgg aac gtg cta gca cag gag      634
Leu Tyr Val Lys Asp Pro Ala Arg Pro Trp Asn Val Leu Ala Gln Glu
      100         105         110

gtg gtc gtg ttc gag gac cag gac gca cta ctg ccc tgt ctg ctc aca      682
Val Val Val Phe Glu Asp Gln Asp Ala Leu Leu Pro Cys Leu Leu Thr
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135 140 145	
 ccc ctc atg cgc cac acc aac tac tcc ttc tcg ccc tgg cat ggc ttc <sup>778</sup>	
Pro Leu Met Arg His Thr Asn Tyr Ser Phe Ser Pro Trp His Gly Phe	
150 155 160	
 acc atc cac agg gcc aag ttc att cag agc cag gac tat caa tgc agt <sup>826</sup>	
Thr Ile His Arg Ala Lys Phe Ile Gln Ser Gln Asp Tyr Gln Cys Ser	
165 170 175	
 gcc ctg atg ggt ggc agg aag gtg atg tcc atc agc atc cgg ctg aaa <sup>874</sup>	
Ala Leu Met Gly Gly Arg Lys Val Met Ser Ile Ser Ile Arg Leu Lys	
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 gtg cag aaa gtc atc cca ggg ccc cca gcc ttg aca ctg gtg cct gca <sup>922</sup>	
Val Gln Lys Val Ile Pro Gly Pro Ala Leu Thr Leu Val Pro Ala	
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Glu Leu Val Arg Ile Arg Gly Glu Ala Ala Gln Ile Val Cys Ser Ala	
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 agc agc gtt gtt aac ttt gat gtc ttc ctc caa cac aac aac acc <sup>1018</sup>	
Ser Ser Val Asp Val Asn Phe Asp Val Phe Leu Gln His Asn Asn Thr	
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 aag ctc gca atc cct caa caa tct gac ttt cat aat aac cgt tac caa <sup>1066</sup>	
Lys Leu Ala Ile Pro Gln Gln Ser Asp Phe His Asn Asn Arg Tyr Gln	
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 aaa gtc ctg acc ctc aac ctc gat caa gta gat ttc caa cat gcc ggc <sup>1114</sup>	
Lys Val Leu Thr Leu Asn Leu Asp Gln Val Asp Phe Gln His Ala Gly	
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 aac tac tcc tgc gtg gcc agc aac gtg cag ggc aag cac tcc acc tcc <sup>1162</sup>	
Asn Tyr Ser Cys Val Ala Ser Asn Val Gln Gly Lys His Ser Thr Ser	
275 280 285 290	
 atg ttc ttc cgg gtg gta gag agt gcc tac ttg aac ttg agc tct gag <sup>1210</sup>	
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295 300 305	
 cag aac ctc atc cag gag gtg acc gtg ggg gag ggg ctc aac ctc aaa <sup>1258</sup>	
Gln Asn Leu Ile Gln Glu Val Thr Val Gly Glu Gly Leu Asn Leu Lys	
310 315 320	
 gtc atg gtg gag gcc tac cca ggc ctg caa ggt ttt aac tgg acc tac <sup>1306</sup>	
Val Met Val Glu Ala Tyr Pro Gly Leu Gln Gly Phe Asn Trp Thr Tyr	
325 330 335	
 ctg gga ccc ttt tct gac cac cag cct gag ccc aag ctt gct aat gct <sup>1354</sup>	
Leu Gly Pro Phe Ser Asp His Gln Pro Glu Pro Lys Leu Ala Asn Ala	
340 345 350	
 acc acc aag gac aca tac agg cac acc ttc acc ctc tct ctg ccc cgc <sup>1402</sup>	
Thr Thr Lys Asp Thr Tyr Arg His Thr Phe Thr Leu Ser Leu Pro Arg	
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 ctg aag ccc tct gag gct ggc cgc tac tcc ttc ctg gcc aga aac cca <sup>1450</sup>	
Leu Lys Pro Ser Glu Ala Gly Arg Tyr Ser Phe Leu Ala Arg Asn Pro	

375	380	385	
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gat gac cca tac cct gag gtc ctg agc cag gag ccc ttc cac aag gtg Asp Asp Pro Tyr Pro Glu Val Leu Ser Gln Glu Pro Phe His Lys Val 455 460 465			1690
acg gtg cag agc ctg ctg act gtt gag acc tta gag cac aac caa acc Thr Val Gln Ser Leu Leu Thr Val Glu Thr Leu Glu His Asn Gln Thr 470 475 480			1738
tac gag tgc agg gcc cac aac agc gtg ggg agt ggc tcc tgg gcc ttc Tyr Glu Cys Arg Ala His Asn Ser Val Gly Ser Gly Ser Trp Ala Phe 485 490 495			1786
ata ccc atc tct gca gga gcc cac acg cat ccc ccg gat gag ttc ctc Ile Pro Ile Ser Ala Gly Ala His Thr His Pro Pro Asp Glu Phe Leu 500 505 510			1834
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ctg ctg ctc ctg cta ttg tac aag tat aag cag aag ccc aag tac Leu Leu Leu Leu Leu Leu Tyr Lys Tyr Lys Gln Lys Pro Lys Tyr 535 540 545			1930
cag gtc cgc tgg aag atc atc gag agc tat gag ggc aac agt tat act Gln Val Arg Trp Lys Ile Ile Glu Ser Tyr Glu Gly Asn Ser Tyr Thr 550 555 560			1978
ttc atc gac ccc acg cag ctg cct tac aac gag aag tgg gag ttc ccc Phe Ile Asp Pro Thr Gln Leu Pro Tyr Asn Glu Lys Trp Glu Phe Pro 565 570 575			2026
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aag gtg gtg gag gcc acg gcc ttt ggt ctg ggc aag gag gat gct gtc Lys Val Val Glu Ala Thr Ala Phe Gly Leu Gly Lys Glu Asp Ala Val 595 600 605 610			2122
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Lys Glu Ala Leu Met Ser Glu Leu Lys Ile Met Ser His Leu Gly Gln			
630	635	640	
cac gag aac atc gtc aac ctt ctg gga gcc tgt acc cat gga ggc cct			2266
His Glu Asn Ile Val Asn Leu Leu Gly Ala Cys Thr His Gly Gly Pro			
645	650	655	
gta ctg gtc atc acg gag tac tgt tgc tat ggc gac ctg ctc aac ttt			2314
Val Leu Val Ile Thr Glu Tyr Cys Cys Tyr Gly Asp Leu Leu Asn Phe			
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ctg cga agg aag gct gag gcc atg ctg gga ccc agc ctg agc ccc ggc			2362
Leu Arg Arg Lys Ala Glu Ala Met Leu Gly Pro Ser Leu Ser Pro Gly			
675	680	685	690
cag gac ccc gag gga ggc gtc gac tat aag aac atc cac ctc gag aag			2410
Gln Asp Pro Glu Gly Val Asp Tyr Lys Asn Ile His Leu Glu Lys			
695	700	705	
aaa tat gtc cgc agg gac agt ggc ttc tcc agc cag ggt gtg gac acc			2458
Lys Tyr Val Arg Arg Asp Ser Gly Phe Ser Ser Gln Gly Val Asp Thr			
710	715	720	
tat gtg gag atg agg cct gtc tcc act tct tca aat gac tcc ttc tct			2506
Tyr Val Glu Met Arg Pro Val Ser Thr Ser Ser Asn Asp Ser Phe Ser			
725	730	735	
gag caa gac ctg gac aag gag gat gga cgg ccc ctg gag ctc cgg gac			2554
Glu Gln Asp Leu Asp Lys Glu Asp Gly Arg Pro Leu Glu Leu Arg Asp			
740	745	750	
ctg ctt cac ttc tcc agc caa gta gcc cag ggc atg gcc ttc ctc gct			2602
Leu Leu His Phe Ser Ser Gln Val Ala Gln Gly Met Ala Phe Leu Ala			
755	760	765	770
tcc aag aat tgc atc cac cgg gac gtg gca gcg cgt aac gtg ctg ttg			2650
Ser Lys Asn Cys Ile His Arg Asp Val Ala Ala Arg Asn Val Leu Leu			
775	780	785	
acc aat ggt cat gtg gcc aag att ggg gac ttc ggg ctg gct agg gac			2698
Thr Asn Gly His Val Ala Lys Ile Gly Asp Phe Gly Leu Ala Arg Asp			
790	795	800	
atc atg aat gac tcc aac tac att gtc aag ggc aat gcc cgc ctg cct			2746
Ile Met Asn Asp Ser Asn Tyr Ile Val Lys Gly Asn Ala Arg Leu Pro			
805	810	815	
gtg aag tgg atg gcc cca gag agc atc ttt gac tgt gtc tac acg gtt			2794
Val Lys Trp Met Ala Pro Glu Ser Ile Phe Asp Cys Val Tyr Thr Val			
820	825	830	
cag agc gac gtc tgg tcc tat ggc atc ctc ctc tgg gag atc ttc tca			2842
Gln Ser Asp Val Trp Ser Tyr Gly Ile Leu Leu Trp Glu Ile Phe Ser			
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ctt ggg ctg aat ccc tac cct ggc atc ctg gtg aac agc aag ttc tat			2890
Leu Gly Leu Asn Pro Tyr Pro Gly Ile Leu Val Asn Ser Lys Phe Tyr			
855	860	865	
aaa ctg gtg aag gat gga tac caa atg gcc cag cct gca ttt gcc cca			2938
Lys Leu Val Lys Asp Gly Tyr Gln Met Ala Gln Pro Ala Phe Ala Pro			
870	875	880	

aag aat ata tac agc atc atg cag gcc tgc tgg gcc ttg gag ccc acc Lys Asn Ile Tyr Ser Ile Met Gln Ala Cys Trp Ala Leu Glu Pro Thr 885 890 895	2986
cac aga ccc acc ttc cag cag atc tgc tcc ttc ctt cag gag cag gcc His Arg Pro Thr Phe Gln Gln Ile Cys Ser Phe Leu Gln Glu Gln Ala 900 905 910	3034
caa gag gac agg aga gag cgg gac tat acc aat ctg ccg agc agc agc Gln Glu Asp Arg Arg Glu Arg Asp Tyr Thr Asn Leu Pro Ser Ser Ser 915 920 925 930	3082
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Lys Pro Gly Ala Thr Val Thr Leu Arg Cys Val Gly Asn Gly Ser Val  
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Glu Trp Asp Gly Pro Pro Ser Pro His Trp Thr Leu Tyr Ser Asp Gly  
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Ser Ser Ser Ile Leu Ser Thr Asn Asn Ala Thr Phe Gln Asn Thr Gly  
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Thr Tyr Arg Cys Thr Glu Pro Gly Asp Pro Leu Gly Gly Ser Ala Ala  
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Ile His Leu Tyr Val Lys Asp Pro Ala Arg Pro Trp Asn Val Leu Ala  
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Gln Glu Val Val Val Phe Glu Asp Gln Asp Ala Leu Leu Pro Cys Leu  
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Leu Thr Asp Pro Val Leu Glu Ala Gly Val Ser Leu Val Arg Val Arg  
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Gly Arg Pro Leu Met Arg His Thr Asn Tyr Ser Phe Ser Pro Trp His  
 145                  150                  155                  160

Gly Phe Thr Ile His Arg Ala Lys Phe Ile Gln Ser Gln Asp Tyr Gln  
 165                  170                  175

Cys Ser Ala Leu Met Gly Gly Arg Lys Val Met Ser Ile Ser Ile Arg  
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Leu Lys Val Gln Lys Val Ile Pro Gly Pro Pro Ala Leu Thr Leu Val  
 195                  200                  205

Pro Ala Glu Leu Val Arg Ile Arg Gly Glu Ala Ala Gln Ile Val Cys  
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Ser Ala Ser Ser Val Asp Val Asn Phe Asp Val Phe Leu Gln His Asn  
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Asn Thr Lys Leu Ala Ile Pro Gln Gln Ser Asp Phe His Asn Asn Arg  
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Tyr Gln Lys Val Leu Thr Leu Asn Leu Asp Gln Val Asp Phe Gln His  
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Ala Gly Asn Tyr Ser Cys Val Ala Ser Asn Val Gln Gly Lys His Ser  
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Ser Glu Gln Asn Leu Ile Gln Glu Val Thr Val Gly Glu Gly Leu Asn  
305 310 315 320

Leu Lys Val Met Val Glu Ala Tyr Pro Gly Leu Gln Gly Phe Asn Trp  
325 330 335

Thr Tyr Leu Gly Pro Phe Ser Asp His Gln Pro Glu Pro Lys Leu Ala  
340 345 350

Asn Ala Thr Thr Lys Asp Thr Tyr Arg His Thr Phe Thr Leu Ser Leu  
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Pro Arg Leu Lys Pro Ser Glu Ala Gly Arg Tyr Ser Phe Leu Ala Arg  
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Asn Pro Gly Gly Trp Arg Ala Leu Thr Phe Glu Leu Thr Leu Arg Tyr  
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Pro Pro Glu Val Ser Val Ile Trp Thr Phe Ile Asn Gly Ser Gly Thr  
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Gln Cys Ser Gly His Thr Asp Arg Cys Asp Glu Ala Gln Val Leu Gln  
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Val Trp Asp Asp Pro Tyr Pro Glu Val Leu Ser Gln Glu Pro Phe His  
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Lys Val Thr Val Gln Ser Leu Leu Thr Val Glu Thr Leu Glu His Asn  
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Gln Thr Tyr Glu Cys Arg Ala His Asn Ser Val Gly Ser Gly Ser Trp  
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Ala Phe Ile Pro Ile Ser Ala Gly Ala His Thr His Pro Pro Asp Glu  
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Phe Leu Phe Thr Pro Val Val Ala Cys Met Ser Ile Met Ala Leu  
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Tyr Thr Phe Ile Asp Pro Thr Gln Leu Pro Tyr Asn Glu Lys Trp Glu  
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Phe Gly Lys Val Val Glu Ala Thr Ala Phe Gly Leu Gly Lys Glu Asp  
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Ala Val Leu Lys Val Ala Val Lys Met Leu Lys Ser Thr Ala His Ala  
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Gly Gln His Glu Asn Ile Val Asn Leu Leu Gly Ala Cys Thr His Gly  
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Gly Pro Val Leu Val Ile Thr Glu Tyr Cys Cys Tyr Gly Asp Leu Leu  
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Asn Phe Leu Arg Arg Lys Ala Glu Ala Met Leu Gly Pro Ser Leu Ser  
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Pro Gly Gln Asp Pro Glu Gly Val Asp Tyr Lys Asn Ile His Leu  
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Glu Lys Lys Tyr Val Arg Arg Asp Ser Gly Phe Ser Ser Gln Gly Val  
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Asp Thr Tyr Val Glu Met Arg Pro Val Ser Thr Ser Ser Asn Asp Ser  
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Phe Ser Glu Gln Asp Leu Asp Lys Glu Asp Gly Arg Pro Leu Glu Leu  
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Arg Asp Leu Leu His Phe Ser Ser Gln Val Ala Gln Gly Met Ala Phe  
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Leu Ala Ser Lys Asn Cys Ile His Arg Asp Val Ala Ala Arg Asn Val  
770 775 780

Leu Leu Thr Asn Gly His Val Ala Lys Ile Gly Asp Phe Gly Leu Ala  
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Arg Asp Ile Met Asn Asp Ser Asn Tyr Ile Val Lys Gly Asn Ala Arg  
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Leu Pro Val Lys Trp Met Ala Pro Glu Ser Ile Phe Asp Cys Val Tyr  
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Thr Val Gln Ser Asp Val Trp Ser Tyr Gly Ile Leu Leu Trp Glu Ile  
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Phe Ser Leu Gly Leu Asn Pro Tyr Pro Gly Ile Leu Val Asn Ser Lys  
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Ala Pro Lys Asn Ile Tyr Ser Ile Met Gln Ala Cys Trp Ala Leu Glu  
885 890 895

Pro Thr His Arg Pro Thr Phe Gln Gln Ile Cys Ser Phe Leu Gln Glu  
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Gln Ala Gln Glu Asp Arg Arg Glu Arg Asp Tyr Thr Asn Leu Pro Ser  
915 920 925

Ser Ser Arg Ser Gly Gly Ser Gly Ser Ser Ser Glu Leu Glu Glu  
930 935 940

Glu Ser Ser Ser Glu His Leu Thr Cys Cys Glu Gln Gly Asp Ile Ala  
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Gln Pro Leu Leu Gln Pro Asn Asn Tyr Gln Phe Cys  
965 970